Tumor Type	Cell Line	Kyriazis (19)	AntiCancer MetaMouse	Clinical Pattern of Metastasis
Bladder (Transitional cell carcinoma)	RT-4	No metastasis	Liver, pancreas, diaphragm, omentum, iliac lymph nodes, superficial inguinal lymph nodes, gastric lymph nodes (1).	Regional lymph nodes, liver, lung, pancreas, diaphragm, spleen (2, Chapter 107).
Bladder (Transitional cell carcinoma)	RT-10		Liver, lung, pancreas, spleen, diaphragm, lymph nodes (21).	
Bladder (Transitional cell carcinoma)	SW-800	Submaxillary lymph node, salivary gland, diaphragm		
Bladder (Transitional cell carcinoma)	SW-780	Pectoral and intercostal muscles, mediastinal lymph nodes.		1
Bladder (Mucinous adenocarcinoma)	13678	Inconsistent finding		

Exhibit 2 RMH

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Exhibit A

Tumor Type	Cell Line	Kyriazis (19)	AntiCancer MetaMouse	Clinical Pattern of Metastasis
Colon	SW-480 (Well differentiated adenocarcinoma)	Lymph node and lungs		Liver, mesenteric lymph nodes, omentum, peritoneum, lung, abdominal wall, disseminated
Colon	Co-3 (Well differentiated adenocarcinoma)		Liver, peritoneum, mesenteric lymph nodes, lung, omentum, abdominal wall, ileum (6-8).	carcinomatosis (2, Chapter 103).
Colon	AC-1935 (Moderately differentiated adenocarcinoma)		Liver, peritoneum, mesenteric lymph nodes, lung, abdominal wall (9).	
Colon	COL-2-JCK (Poorly differentiated adenocarcinoma)		Liver (10,11).	·
Colon	COL-3-JCK (Poorly differentiated adenocarcinoma)		Liver (11). Mesenteric lymph nodes (7).	
Colon	COL-5-JCK (Well differentiated adenocarcinoma)		Liver (11).	·
Colon	Patient colon cancer specimens		Liver, lymph nodes, local invasion, disseminated carcinomatosis (12).	

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Tumor Type	Cell Line	Kyriazis (19)	AntiCancer MetaMouse	Clinical Pattern of Metastasis
Pancreas	(Mia)PaCa (Well differentiated adenocarcinoma)	Axillary lymph nodes, lung.	Liver, spleen, portal lymph nodes, stomach, mediastinum, lung, retroperitoneum (13, 16, 20).	Liver, spleen, portal lymph nodes, colon, stomach, mediastinum, lung, kidney, retroperitoneum, diaphragm, small intestine (2,
Pancreas	Capan-1	Lymph nodes, lung.		Chapter 101).
Pancreas	Patient specimens		Liver, stomach, duodenum, regional lymph nodes, adrenal gland, diaphragm, mediastinal lymph nodes (14).	
Pancreas	Panc-4		Liver, peritoneal, duodenum (15).	
Pancreas	BxPC-3		Portal lymph nodes, retroperitoneum, spleen, liver, diaphragm, small intestine, colon, liver, kidney, mediastinum, lung, omentum (13, 16,20).	
Pancreas	Pan-12-JCK		Liver, kidney, regional and distant lymph nodes, lung, adrenal gland (17,18).	

Tumor Type	Cell Line	Kyriazis (19)	AntiCancer MetaMouse	Clinical Pattern of Metastasis
Breast	BrCa	Lymph nodes		Axillary lymph nodes, lung, liver,
Breast	MDA-MB-435		Axillary lymph nodes, lung, liver (3). Bone (4).	bone (2, Chapter 118).
Breast	AC-2468 (Patient breast		Lung (5).	

REFERENCES:

- Fu, X., and Hoffman, R.M. Human RT-4 bladder carcinoma is highly metastatic in nude mice and comparable to ras-H-transformed RT-4 when orthotopically onplanted as histologically intact tissue. Int. J. Cancer 51, 989-991, 1992.
- Holland, James E., and Frei, Emil III (eds.), Cancer Medicine, 5th Edition. Hamilton, Ontario, Canada: B.C. Decker, Inc., 2000. α
- Li, X-M., Wang, J-W., An, Z., Yang, M. Baranov, E., Jiang, P., Sun, F-X., Moossa, A.R., and Hoffman, R.M. Optically-imageable metastatic model of human breast cancer. Clinical & Experimental Metastasis, in press. 3
- 4. An, Z., et al., unpublished.
- Fu, X., Le, P., and Hoffman, R.M. A metastatic orthotopic-transplant nude-mouse model of human patient breast cancer. Anticancer Res. 13, 901-904, Ś
- Fu, X., Herrera, H., Kubota, T., and Hoffman, R.M. Extensive liver metastasis from human colon cancer in nude and scid mice after orthotopic onplantation of histologically-intact human colon carcinoma tissue. Anticancer Res. 12, 1395-1398, 1992. છં
- Togo, S., Shimada, H., Kubota, T., Moossa, A.R., Hoffman, R.M. Host organ specifically determines cancer progression. Cancer Res. 55, 681-684, 1995. 7
- An, Z., Wang, X., Willmott, N., Chander, S.K., Tickle, S., Docherty, A.J.P., Mountain, A., Millican, A.T., Morphy, R., Porter, J.R., Epemolu, R.O., Kubota, T., Moossa, A.R., and Hoffman, R.M. Conversion of highly malignant colon cancer from an aggesssive to a controlled disease by oral administration of a metalloproteinase inhibitor. Clinical & Experimental Metastasis 15, 184-195, 1997. ∞i
- Wang, X., Fu, X., Brown, P.D., Crimmin, M.J., and Hoffman, R.M. Matrix metalloproteinase inhibitor BB-94 (Batimastat) inhibits human colon tumor growth and spread in a patient-like orthotopic model in nude mice. Cancer Res. 54, 4726-4728, 1994. 6
- Immunochemotherapy prevents human colon cancer metastasis after orthotopic onplantation of histologically-intact tumor tissue in nude mice. Anticancer Furukawa, T., Kubota, T., Watanabe, M., Kuo, P.H., Kase, S., Saikawa, Y., Tanino, H., Teramoto, T., Ishibiki, K., Kitajima, M., and Hoffman, R.M. Res. 13, 287-291, 1993. <u>.</u>
- Kuo, T-H., Kubota, T., Watanbe, M., Furukawa, T., Teramoto, T., Ishibiki, K., Kitajimi, M., Moossa, A.R., Penman, S., Hoffman, R.M. Liver colonization competence governs colon cancer metastasis. Proc. Natl. Acad. Sci. USA 92, 12085-12089, 1995. Ξ
- Fu, X., Besterman, J.M., Monosov, A., and Hoffman, R.M. Models of human metastatic colon cancer in nude mice orthotopically constructed by using nistologically intact patient specimens. Proc. Natl. Acad. Sci. USA 88, 9345-9349, 1991. 2

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- Bouvet, M., Yang, M., Nardin, S., Wang, X., Jiang, P., Baranov, E., Moossa, A.R., Hoffman, R.M. Chronologically-specific metastatic targeting of numan pancreatic tumors in orthotopic models. Clinical & Experimental Metastasis 18, 213-218, 2000. <u>.</u>
- Fu, X., Guadagni, F., and Hoffman, R.M. A metastatic nude-mouse model of human pancreatic cancer constructed orthotopically from histologically ntact patient specimens. Proc. Natl. Acad. Sci. USA 89, 5645-5649, 1992. 7.
- Furukawa, T., Kubota, T., Watanabe, M., Kitajima, M., and Hoffman, R.M. A novel "patient-like" treatment model of human pancreatic cancer constructed using orthotopic transplantation of histologically intact human tumor tissue in nude mice. Cancer Res. 53, 3070-3072, 1993. 15.
- Lee, N.C., Bouvet, M., Nardin, S., Jiang, P., Baranov, E., Rashidi, B., Yang, M., Wang, X., Moossa, A.R., and Hoffman, R.M. Antimetastatic efficacy of adjuvant gemcitabine in a pancreatic cancer orthotopic model. Clinical & Experimental Metastasis 18, 379-384, 2001. 16.
- An, Z., Wang, X., Kubota, T., Moossa, A.R., Hoffman, R.M. A clinical nude mouse metastatic model for highly malignant human pancreatic cancer. Anticancer Res. 16, 627-632, 1996. 17.
- Tomikawa, M., Kubota, T., Matsuzaki, S.W., Takahasi, S., Kitajima, M., Moossa, A.R., and Hoffman, R.M. Mitomycin C and cisplatin increase survival in a human pancreatic cancer metastatic model. Anticancer Res. 17, 3623-3626, 1997. ∞.
- Kyriazis, A.P., Kyriazis, A.A., McCombs, William B. III, and Kereiakes, J.A. Biological behavior of human malignant tumors grown in the nude mouse. Cancer Res. 41, 3995-4000, 1981. 19
- Bouvet, M., Wang, J-W., Nardin, S.R., Nassirpour, R., Yang, M., Baranov, E., Jiang, P., Moossa, A.R., and Hoffman, R.M. Real-time optical imaging of primary tumor growth and multiple metastatic events in a pancreatic cancer orthotopic model. Cancer Research 62, 1534-1540, 2002. 20.
- Fu, X., Theodorescu, D., Kerbel, R.S., and Hoffman, R.M. Extensive multi-organ metastasis following orthotopic onplantation of histologically-intact human bladder carcinoma tissue in nude mice. Int. J. Cancer 49, 938-939, 1991. 21.

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Tumor Type	Cell Line	Otto (1)	AntiCancer MetaMouse	Clinical Pattern of Metastasis
Renal cell carcinoma	RCC 7 RCC 9 RCC 14	No metastasis.		Lung, lymph nodes, liver, and brain (3, Chapter 1057).
	SN12C		Lung, lymph nodes, liver.	

REFERENCES:

- Otto, U., Huland, H., Baisch, H., and Kloppel, G. Transplantation of human renal cell carcinoma into NMRI nu/nu mice. III. Effect of irradiation on tumor acceptance and tumor growth. J. Urol. 134, 170-174, 1985.
- carcinoma in nude mice: benefits of fragment implantation compared to cell-suspension injection. Clinical & Experimental Metastasis 17, An, Z., Jiang, P., Wang, X., Moossa, A.R., and Joffman, R.M. Development of a high metastatic orthotopic model of human renal cell 265-270, 1999. \sim i
- Holland, James E., and Frei, Emil III (eds.), Cancer Medicine, 5th Edition. Hamilton, Ontario, Canada: B.C. Decker, Inc., 2000. ω

Exhibit 3 RMH